

AMENDMENTS TO THE CLAIMS: This listing of claims replaces all prior versions and listings of claims in this patent application.

1-119. (canceled)

120. (currently amended) A composition comprising a duplex consisting of a first chemically synthesized oligomeric compound and a second chemically synthesized oligomeric compound, wherein:

each of the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound independently consists of 17 to 25 linked nucleosides, each nucleoside comprising a nucleobase and a sugar;

~~at least 17 contiguous nucleobases of~~ the first chemically synthesized oligomeric compound ~~are~~ is 100% complementary to ~~at least 17 contiguous nucleobases of~~ the second chemically synthesized oligomeric compound and to a target mRNA;

the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound are not covalently linked to each other; and

the first chemically synthesized oligomeric compound is a gapmer, wherein the gap comprises at least 4 nucleosides, each comprising a 2'-OH, and wherein each nucleoside of each wing comprises a 2' modification or is an LNA nucleoside.

121. (previously presented) The composition of claim 120, wherein the second chemically synthesized oligomeric compound is a gapmer, wherein the gap comprises at least 4 nucleosides, each comprising a 2'-OH, and wherein each nucleoside of each wing comprises a 2' modification or is an LNA nucleoside.

122-123. (canceled)

124. (previously presented) The composition of claim 120, wherein each nucleoside of at least one of the wings of the gapmer comprises a 2' sugar modification selected from fluoro, alkoxy, amino-alkoxy, allyloxy, imidazolylalkoxy, and methoxyethoxy or is an LNA nucleoside.

125. (withdrawn) The composition of claim 120, wherein each nucleoside of one wing of the gapmer comprises a 2'-fluoro and each nucleoside of the other wing of the gapmer comprises a 2'-OCH₃.

126. (withdrawn) The composition of claim 120, wherin each nucleoside of the 5' wing of the gapmer comprises a 2'-fluoro.

127. (previously presented) The composition of claim 120, wherein each nucleoside of the 3' wing of the gapmer comprises a 2'- OCH₃.

128. (withdrawn) The composition of claim 126, wherein each nucleoside of the 3' wing of the gapmer comprises a 2'- OCH₃.

129-130. (canceled)

131. (withdrawn) The composition of claim 120, wherein each nucleoside of one wing of the gapmer comprises a 2'-fluoro and each nucleoside of the other wing of the gapmer is an LNA.

132. (withdrawn) The composition of claim 120, wherein each nucleoside of the 3' wing of the gapmer is an LNA.

133. (withdrawn) The composition of claim 126, wherein each nucleoside of the 3' wing of the gapmer is an LNA.

134-135. (canceled)

136. (previously presented) The composition of claim 120, wherein each wing of the gapmer is from two to seven nucleosides in length.

137. (previously presented) The composition of claim 120, wherein at least one of the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound comprises at least one phosphorothioate linkage.

138. (previously presented) The composition of claim 137, wherein each of the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound comprises at least one phosphorothioate linkage.

139. (previously presented) A compound comprising a chemically synthesized oligomeric compound, wherein:

the chemically synthesized oligomeric compound consists of 17 to 25 linked nucleosides, each nucleoside comprising a nucleobase and a sugar;

the chemically synthesized oligomeric compound is a gapmer, wherein the gap comprises at least 4 nucleosides, each comprising a 2'-OH, and wherein each nucleoside of each wing comprises a 2' modification or is an LNA nucleoside; and

the chemically synthesized oligomeric compound is an antisense compound.

140. (previously presented) The compound of claim 139, wherein each nucleoside of at least one of the wings of the gapmer comprises a 2' sugar modification selected from fluoro, alkoxy, amino-alkoxy, allyloxy, imidazolylalkoxy, and methoxyethoxy or is an LNA nucleoside.

141. (withdrawn) The compound of claim 139, wherein each nucleoside of one wing of the gapmer comprises a 2'-fluoro and each nucleoside of the other wing of the gapmer comprises a 2'-OCH₃.

142. (withdrawn) The compound of claim 139, wherein each nucleoside of the 5' wing of the gapmer comprises a 2'-fluoro.

143. (previously presented) The compound of claim 139, wherein each nucleoside of the 3' wing of the gapmer comprises a 2'- OCH₃.

144. (withdrawn) The compound of claim 142, wherein each nucleoside of the 3' wing of the gapmer comprises a 2'-OCH₃.

145. (withdrawn) The compound of claim 139, wherein each nucleoside of one wing of the gapmer comprises a 2'-fluoro and each nucleoside of the other wing of the gapmer is an LNA.

146. (withdrawn) The compound of claim 139, wherein each nucleoside of the 3' wing of the gapmer is an LNA.

147. (withdrawn) The compound of claim 142, wherein each nucleoside of the 3' wing of the gapmer is an LNA.

148. (previously presented) The compound of claim 139, wherein each wing of the gapmer is from two to seven nucleosides in length.

149. (previously presented) The compound of claim 139, wherein the chemically synthesized oligomeric compound comprises at least one phosphorothioate linkage.

150. (withdrawn) A composition comprising a duplex consisting of a first chemically synthesized oligomeric compound and a second chemically synthesized oligomeric compound, wherein:

each of the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound independently consists of 17 to 25 linked nucleosides, each nucleoside comprising a nucleobase and a sugar;

at least 17 contiguous nucleobases of the first chemically synthesized oligomeric compound are 100% complementary to at least 17 contiguous nucleobases of the second chemically synthesized oligomeric compound;

the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound are not covalently linked to each other; and

the first chemically synthesized oligomeric compound is a blockmer comprising at least one region comprising at least 4 contiguous nucleosides, each comprising a 2'-OH, and a block region, wherein each nucleoside of the block comprises a 2' modification or is an LNA nucleoside.

151. (withdrawn) The composition of claim 150, wherein the second chemically synthesized oligomeric compound is a blockmer comprising at least one region comprising at least 4 contiguous nucleosides, each comprising a 2'-OH, and a block region, wherein each nucleoside of the block comprises a 2' modification or is an LNA nucleoside.

152. (withdrawn) The composition of claim 150, wherein the duplex comprises an overhang on the 5' end, on the 3' end, or on both the 5' end and the 3' end.

153. (withdrawn) The compound of claim 150, wherein the duplex is blunt-ended.

154. (withdrawn) The composition of claim 150, wherein each nucleoside of the block of the blockmer comprises a 2' sugar modification selected from fluoro, alkoxy, amino-alkoxy, allyloxy, imidazolylalkoxy, and methoxyethoxy or is an LNA nucleoside.

155-156. (canceled)

157. (withdrawn) The composition of claim 150, wherein the block is from two to seven nucleosides in length.

158. (withdrawn) The composition of claim 150, wherein the block is at the 3' terminal end of the chemically synthesized oligomeric compound.

159. (withdrawn) The composition of claim 150, wherein the block is at the 5' terminal end of the chemically synthesized oligomeric compound.

160. (withdrawn) The composition of claim 150, wherein at least one of the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound comprises at least one phosphorothioate linkage.

161. (withdrawn) The composition of claim 150, wherein each of the first chemically synthesized oligomeric compound and the second chemically synthesized oligomeric compound comprises at least one phosphorothioate linkage.

162. (withdrawn) A compound comprising a chemically synthesized oligomeric compound, wherein:

the chemically synthesized oligomeric compound consists of 17 to 25 linked nucleosides, each nucleoside comprising a nucleobase and a sugar;

the chemically synthesized oligomeric compound is a blockmer comprising at least one region comprising at least 4 contiguous nucleosides, each comprising a 2'-OH, and a block region, wherein each nucleoside of the block comprises a 2' modification or is an LNA nucleoside; and

the chemically synthesized oligomeric compound is an antisense compound.

163. (withdrawn) The compound of claim 162, wherein each nucleoside of the block of the blockmer comprises a 2' sugar modification selected from fluoro, alkoxy, amino-alkoxy, allyloxy, imidazolylalkoxy, and methoxyethoxy or is an LNA nucleoside.

164. (withdrawn) The compound of claim 162, wherein the block of the blockmer is from two to seven nucleosides in length.

165. (withdrawn) The compound of claim 162, wherein the block of the blockmer is at the 5' end of the chemically synthesized oligomeric compound.

166. (withdrawn) The compound of claim 162, wherein the block of the blockmer is at the 3' end of the chemically synthesized oligomeric compound.

167. (withdrawn) The compound of claim 162, wherein at least one of the chemically synthesized oligomeric compound comprises at least one phosphorothioate linkage.